



**Aspen Investor Visit
Port Elizabeth Site,
South Africa**

10 May 2011

Aspen's Global Footprint

C = Combined sales, marketing, distribution and manufacturing centres

G = Group headquarters

M = Manufacturing centres

S = Sales, marketing and distribution centres



1. **G** Durban, South Africa
2. **C** Cape Town, South Africa
3. **C** Johannesburg, South Africa
4. **M** Port Elizabeth, South Africa
5. **M** East London, South Africa
6. **M** Toluca, Mexico
7. **S** Mexico City, Mexico
8. **S** Caracas, Venezuela
9. **S** Rio de Janeiro, Brazil
10. **M** Vittoria, Brazil
11. **S** Dublin, Ireland
12. **M** Bad Oldesloe, Germany
13. **C** Nairobi, Kenya
14. **S** Kampala, Uganda
15. **C** Dar es Salaam, Tanzania
16. **S** Dubai, United Arab Emirates
17. **S** Grand Bay, Mauritius
18. **S** Quarry Bay, Hong Kong
19. **C** Sydney, Australia
20. **M** Melbourne, Australia

We remain committed to making a meaningful difference in the health of all those using Aspen medicines globally, and to increase our contribution to the lives of people in the many markets we serve across the globe through the manufacture and supply of high-quality, affordable medicines in a responsible manner.



Aspen's Manufacturing Competitive Advantages

Manufacturing capacity and capability is aligned to the Group's growth strategy

- Aspen's strength lies in its ability to supply high volumes of products reliably with 4.8 billion tablets being manufactured currently
- Manufacturing capabilities at the Port Elizabeth and East London facilities have been rationalised
- Homogenous product types are produced at designated facilities
- Manufacturing capacity in the tableting can be trebled by 7% increase in the expense base with the addition of only incremental variable costs
- Accreditations received from South African and relevant international authorities
- Sufficient manufacturing capacity exists to produce required volumes for the domestic and international markets as well to accommodate the introduction of global brands and Sigma products
- Focus on economies of scale - international volumes are being transferred to Port Elizabeth to reduce reliance on the public sector volumes
- Effective procurement strategies ensure that materials are purchased at competitive prices
- There is a committed focus on manufacturing efficiency projects – benefits have already been realised, and initiatives are already in progress to conserve energy and environmental management

Our world class South African Operations team has created Aspen's sustainably efficient and competitive manufacturing platform

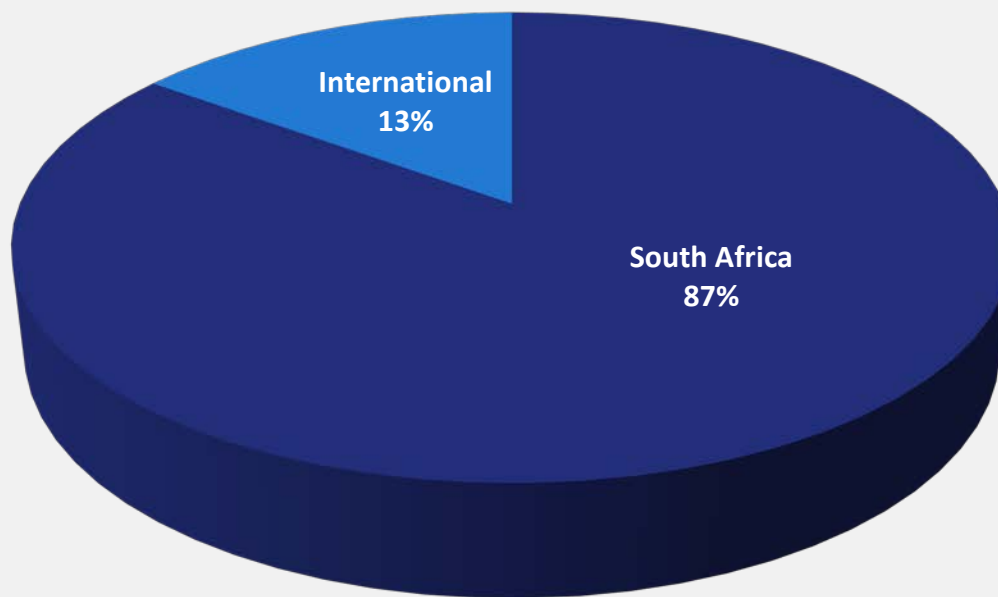
Evolution of Aspen's Manufacturing Base

- Aspen General Facility has been on the present site for approximately 70 years
- Acquired by Aspen from South African Druggists in March 1999, together with facilities in East London and Johannesburg
- Mainly supplied the South African market
- Within 12 years, Aspen has transformed from being a domestically accredited supplier to an international pharmaceutical manufacturer with the developed capability to supply various dosage forms to any pharmaceutical market in the world
- In the last 5 years, more than R2 billion has been invested in the Group's South African facilities for infrastructural expansion and enhancements to improve compliance to the relevant regulatory standards, and in order to support Aspen's sustained supply to both its domestic and diverse international markets
- More than 2700 people are permanently employed at Aspen's Port Elizabeth, East London and Nutritionals manufacturing facilities














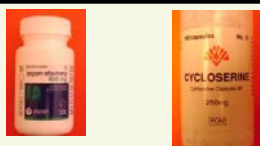






Current Allocation of Domestic and International Volumes

- **Manufactured products are supplied to Europe, Latin America and China**
- **Additional international volumes are in the process of being introduced to the Port Elizabeth facility**
 - ▶ Regulatory approvals are required by the relevant territory, per product, before commercial production can take place



Strategic Manufacturing Partnerships

	Boehringer Ingelheim	Nevirapine	
	GSK	Lamivudine, Zidovudine, Combivir, Epivir & Others	
	BMS	Stavudine, Didanosine, Atazanavir	
	Gilead	Tenofovir & Emtricitabine	
	MSD	Efavirenz	
	Iroko	Aldomet and Indocid	
	Eli Lilly	Cycloserine and Capreomycin	
	Bayer	Nur- Isterate Injection	
	Prestige Brands	Murine & Murine Plus Range of Eye Drop Products	

Aerial view of Aspen's Global Manufacturing Base in Port Elizabeth



Regulatory Authorities Relevant to South African Operations



Regulatory Authority	Unit 1	Unit 2	Unit 3	SVP MP	SVP HP	East London	ADC	ACW
MCC	X	X	X	X	X	X	X	X
FDA	X	#		#	#		X	
MHRA	X	#		#	#		X	
WHO	X	#		#	#		X	
TGA	X	#		#	#		X	
Anvisa (Brazil)	X	#		#	#		X	
	High volume solid manufacturing for domestic and export markets	Small to medium volume solid manufacturing for domestic and export markets: fluid-bed dried products (2A) & oven dried products (2B)	End state solid packing for domestic market	Eye drops, lyophilized vials for domestic and export markets	High potency (incl. hormonal) injectables for domestic and export markets	Semi-solids, liquids and oral contraceptives for domestic market	Warehousing for domestic and export markets	Warehousing for domestic market

x = Approved

= Inspection planned

Strategic Rationalisation Initiatives

Significant efficiencies and cost reductions have been realised through the consolidation of the Port Elizabeth site management systems:

- **Site management and facilities support functions have been consolidated under a single management structure**
 - ▶ Historically two separate entities with individual management structures
 - ▶ Significant rationalisation of redundant costs
- **Consolidation of engineering stores and warehouse management systems**
 - ▶ Integrated site stock management systems and hence focussed working capital management
- **Consolidation of ERP systems for all facilities into a single Baan system**
 - ▶ Provides access to a single reporting platform
 - ▶ Facilitates focussed review, analysis and stratification of site information
 - ▶ Assists with effective optimisation of resources and capability
- **Consolidation of the purchasing systems**
 - ▶ Reduced number of purchase orders
 - ▶ Lower average purchase price per order due to combined volumes
 - ▶ Reduced number of lab tests required on incoming materials due to purchase order rationalisation

Unit 1 – Oral Solid Dosage Facility

- In 2005, an investment was made in the construction of an FDA-approved facility in response to the HIV/AIDS pandemic in Africa
- High volume production of tablets and capsules
- The facility was successfully re-inspected by FDA, MRHA and WHO in 2010
 - ▶ No major findings reported
- Unit 1 currently manufactures 2.8 billion tablets
- A granulation capacity expansion project to the value of R178 million is nearing completion (first suite in operation, with second and third suites in final stages)
 - ▶ This will more double current granulation capacity to increase tableting capacity

Unit 2 – Oral Solid Dosage Facility

Unit 2

- 2.0 billion tablets are manufactured in this area
- Total capacity = 3 billion tablets

Part 2A – Fluid Bed Dried (FBD) products

- Small to medium volume solids production for selected regulated markets
- Reformulated products and investment in new equipment
- Improved process flow and hence more efficient manufacturing process
- Increased automation and improved yields
- Reduction in conversion costs
- 96 tableted products have been transferred into Unit 2A; 40 products will be transferred by July 2011

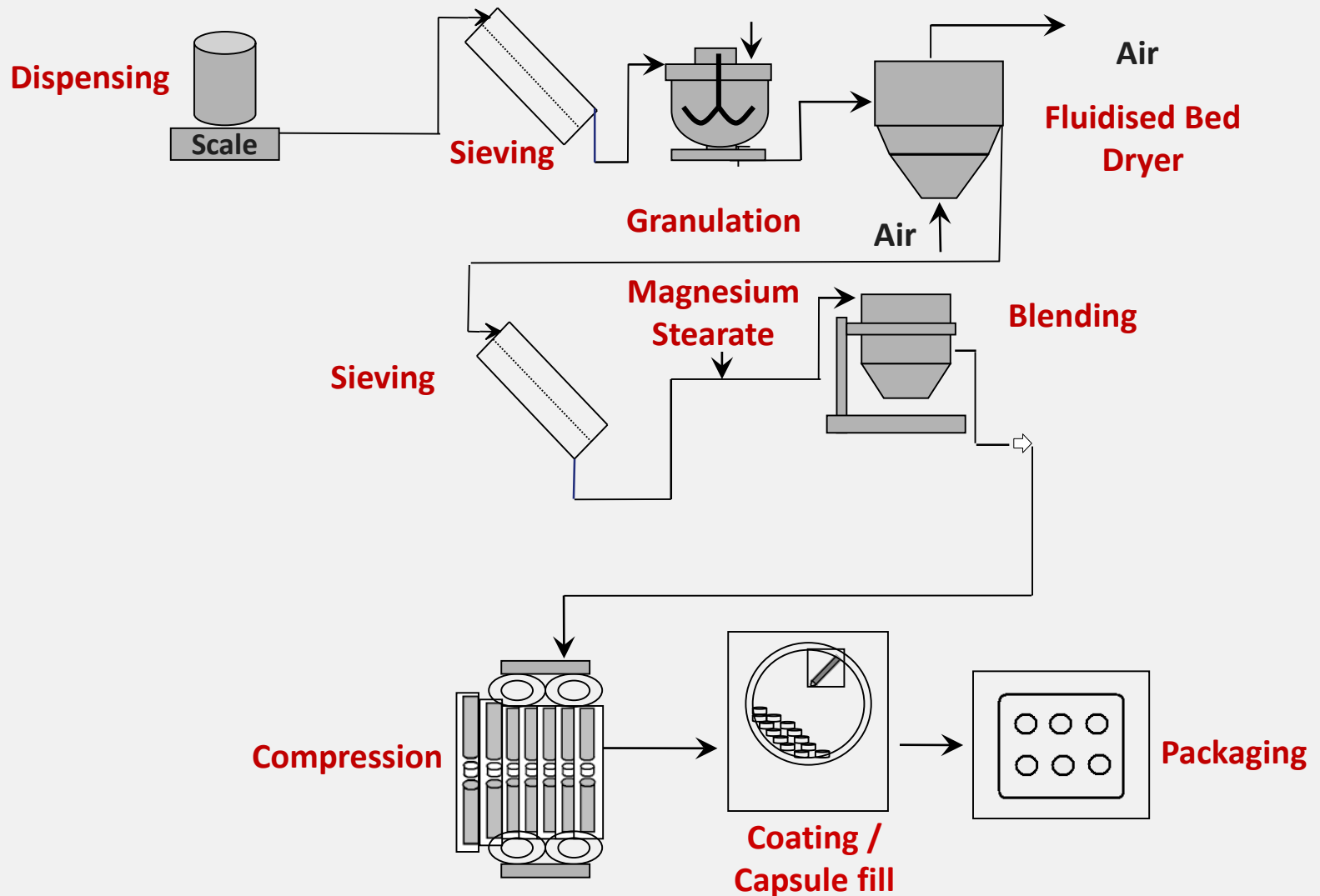
Part 2B – Oven dried products

- Small volume solids production for selected regulated markets
- Investment in new equipment
- Increased automation and improved yields
- 6 products have been transferred into Unit 2B
- 61 products remain to be transferred by April 2012

Unit 3 – General Facility

- A phased process for the transfer of products to Unit 1 and Unit 2 is in progress
- Bulk tablet manufacturing at Unit 3 will be decommissioned by end June 2011
- Semi-solids (creams, ointments and suppositories) and Liquid Lennon Dutch Medicines have been transferred to East London
- Packing for the local market remains in this area

Flow Diagram of the Tableting Processes



Focus on Continuous Improvement Initiatives

Introduction of the Fluid Bed Drying Process

- **The older oven drying technology**
 - ▶ Wet granule obtained during tablet manufacture is dried by spreading materials onto trays inside an oven
 - ▶ Process time is relatively long, requiring an average of 8 - 16 hour drying time
- **The newer FBD technology**
 - ▶ Wet granule obtained during tablet manufacture is sieved into a drying chamber and hot air is passed through the chamber at relatively high speeds, resulting in fluidisation of the granule
 - ▶ Faster process, requiring 2 - 3 hour drying time
- **All key and high volume products have been reformulated and are now manufactured in this area**

Impact: *The benefits of this technology have been extracted and improvements in manufacturing and cost efficiencies have been realised*

Focus on Continuous Improvement Initiatives

- **Doubling of batch sizes to reduce set-up time between batches and thereby optimise equipment utilisation**

- ▶ All Unit 1 manufactured and all Unit 1/3 packed products completed
- ▶ Unit 2 manufactured products in progress

Impact: *Reduction of total annual set up time by 50%*

- **Automation of end-of-line packing to match the increased rate of manufactured output / manual packing processes transferred to automated equipment**

- ▶ Blister and bottle packing completed, and patient ready packing nearing completion
- ▶ Further projects targeted

Impact: *30% Output improvement in patient-ready packs with a reduction in temporary staff*

- **Modification of batch manufacturing records to improve process documentation flow and move to Electronic Batch Documentation has been initiated**

Impact: *Expected reduction in administration time and headcount*

- **Barcode inventory management system introduced in warehousing and now being rolled out to production management**

Impact: *16% Reduction in headcount achieved to date*

Focus on Continuous Improvement Initiatives

- Barcode status labelling of materials implemented to reduce handling activities

Impact: More efficient process through an instantaneous scanning process. Reduction in labelling time - manual capturing and labelling change used take up to one shift

- Evaluation of shared packing tooling in progress

Impact: Will reduce set-up and changeover time significantly to unlock additional capacity

- Re-evaluation of automated (new IBC wash-station) and manual cleaning activities (additional wash-bays at place-of-work and transfer trolleys) in progress

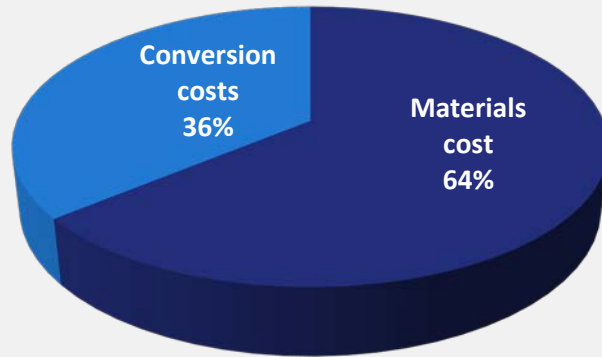
Impact: Will reduce cleaning times to unlock additional capacity

- Evaluation of compression set-up and run-rate parameters proceeding well

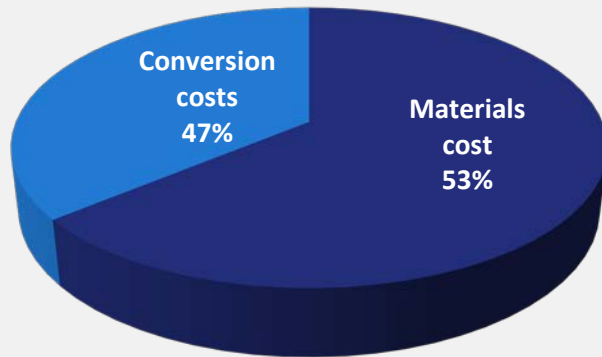
Impact: Increases in run rates of 20% achieved in Unit 1

In-house Manufacturing Cost Analysis

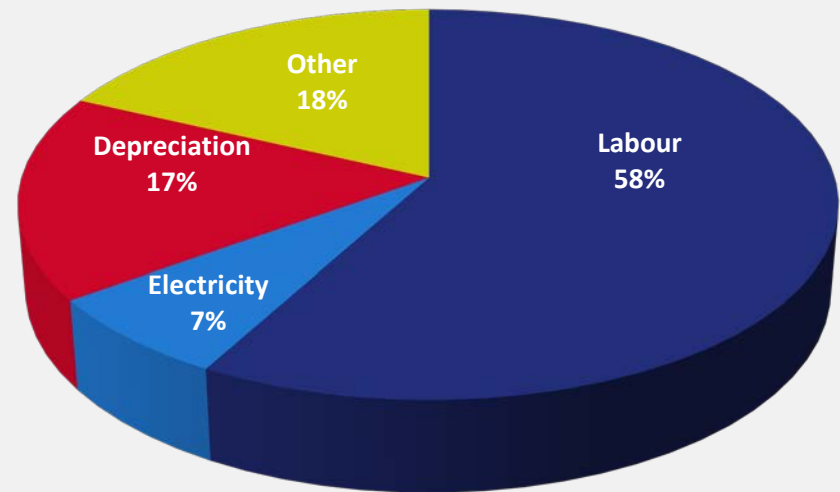
**Allocation of Production Costs
(including ARV's)**



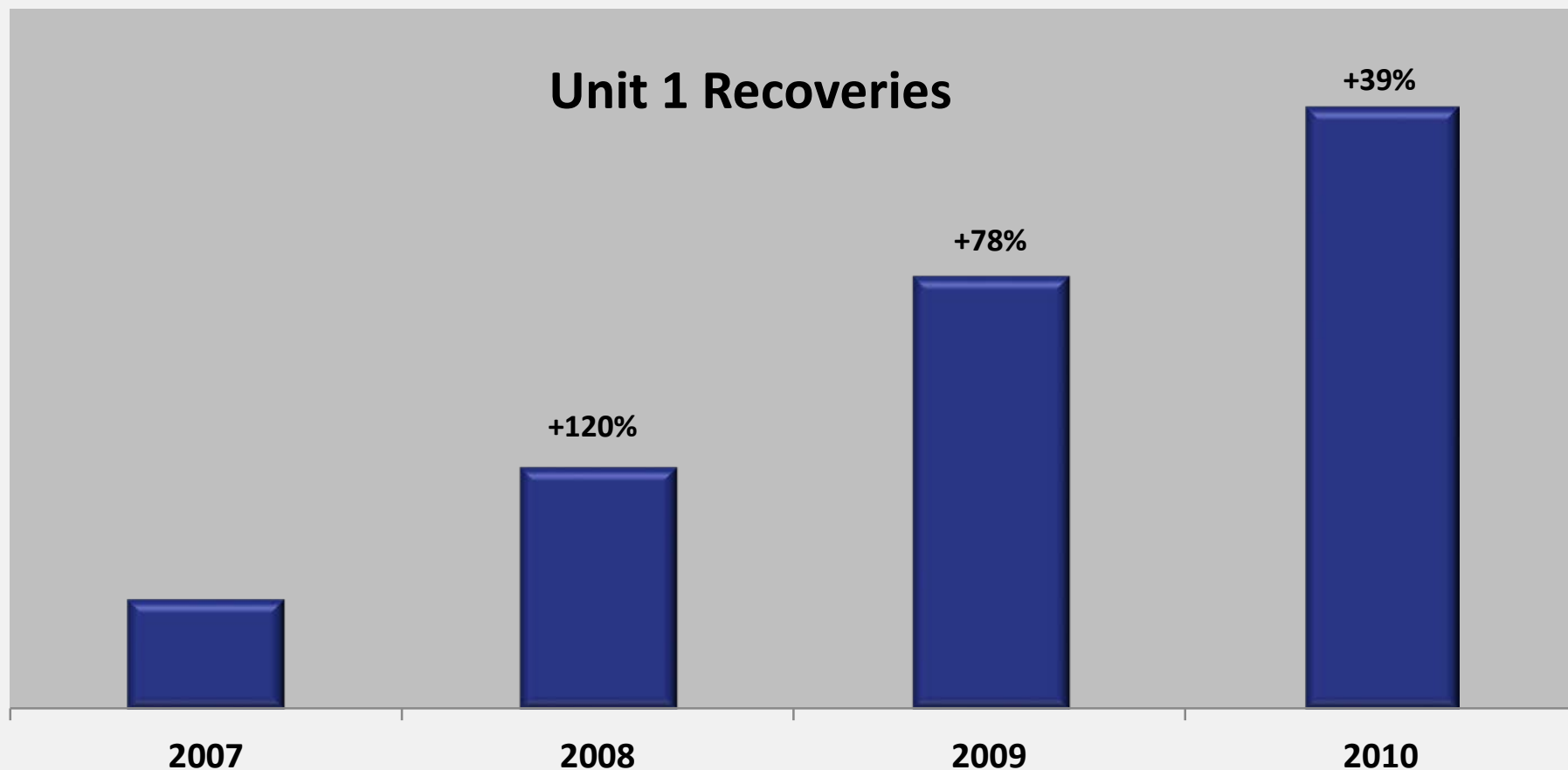
**Allocation of Production Costs
(excluding ARV's)**



Allocation of Conversion costs



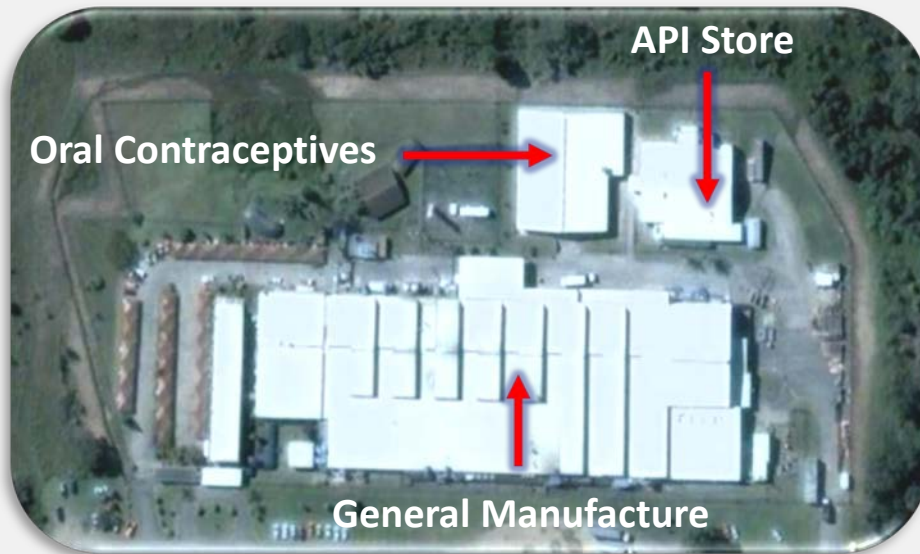
Impact of Continuous Improvement on Recoveries



Prospects in Solids Manufacturing at the Port Elizabeth Site

- **The redundant areas in Unit 3 will be converted into specialised manufacturing areas including**
 - ▶ High potency suite (HPS)
 - ▶ Toxic suite (TS)
- **Following approval of the final design, construction of the HPS and TS will commence in the 2012 year**
 - ▶ High potency and toxic products both require contained manufacturing areas with the air handling and employee health, safety and protective clothing requirements being more stringent in a toxic manufacturing area
- **Approximately 4 billion tablets will be transferred into Unit 1, 2, HPS and TS including**
 - ▶ Selected global brands from Europe
 - ▶ Selected products from Sigma
- **Transfer of these products to Port Elizabeth will almost double manufactured volumes to 8.8 billion tablets (currently 4.8 billion tablets are produced in Unit 1 and 2)**
 - ▶ Economies of scale will be further improved
 - ▶ The additional volumes will be added with only 50% of incremental variable costs being added. The fixed cost base will remain unchanged

East London Site



Aerial view of East London plant



An employee in the Oral Contraceptives facility wearing the necessary protective clothing

East London Multi-Purpose Site

General Facility

- Highly flexible, small volume, niche, solids manufacture, including low dose drugs
- Significant investment has been made into upgrading the East London facility for the manufacture of semi-solids and Dutch medicines

Oral Contraceptive Facility

- The Oral Contraceptive Facility provides a contained manufacturing area for the production of high volume oral contraceptives for the public and private sector

The Steriles Facility

- Project commenced in 2007, presenting challenging complexities of design, technology, technical compliance requirements and specialist skills
- Specialist facility for the manufacture of eye drops, lyophilised vials and sterile injectables, including hormonal vials and ampoules (a niche capability to supply female contraceptives and HRT products)
- The facility commenced with the production of eye drops for the US market in July 2009
- The lyophilised vials area was commercialised in September 2010
- Approved by the MCC for the manufacture of sterile eye drops and lyophilised vials, with WHO approval for Capreomycin being targeted
- Export certificates were granted by the FDA and MCC for the supply of eye drops into the USA
- The sterile facility represents a niche and complex manufacturing

Steriles Facility – Multi-Product Suite

Lines 1 and 2

- High volume eye drops with an annual capacity of 42 million units
- Close to 30 million units being exported to US and Canada under a manufacturing contract with Prestige Brands
- Aspen's Eye Gene range has been introduced; remaining Aspen range of eye drops, i.e. Safyr Bleu and Oculerge in progress

Line 3

- Lyophilised vials have been introduced into this area, starting with Capreomycin, a product for MDR TB
 - ▶ Vancomycin, Clarithromycin and other lyophilised products will be introduced in 2011
- Capacity for lyophilised vials is approximately 2.4 million units
- Additional capacity exists for 2.9 million liquid filled vials
 - ▶ Vitamin B Co, Vitamin B12 and Thiamine liquid filled vials are currently being introduced
- Additional space exists to cater for the introduction of further capabilities - an engineering and design feasibility study has been completed

Steriles Facility – High Potency Suite

Line 4

- High volume ampoule filling capability for hormonal injectables - a niche capability for filling and packing oily solutions which is very relevant to emerging markets
 - ▶ Validation batches of Bayer Nur-Isterate are scheduled for 2011
- Aspen's Lenasone, Betanoid and Decasone are currently being introduced. The MCC have been invited to inspect and approve this area of the facility, where after full commercial production will commence
- Annual capacity for 30 million ampoules

Line 5

- High volume vial filling capability for hormonal injectables - a unique aseptic suspension filling capability
- Aspen's Medroxyprogesterone is envisaged to be manufactured for SA, emerging markets and WHO territories; a global formulation is being developed for world wide supply
- Offtake initially forecast at 20 million vials, annual capacity for vial filling of approximately 45 million units
- Application has recently been made to MCC for inspection of this area in order to facilitate commercial production

Procurement Initiatives

- **Comprehensive understanding of products / materials, driving factors and market factors**
- **Ability to effectively benchmark product prices, quality and supplier service to ensure that we are procuring optimally**
- **Longstanding relationships have been established with international API and packaging suppliers**
- **Ability to source competitively from developing markets**
- **Only reputed and validated suppliers are approved for use**
- **Alternate suppliers are in place to mitigate supply risk and leverage pricing**

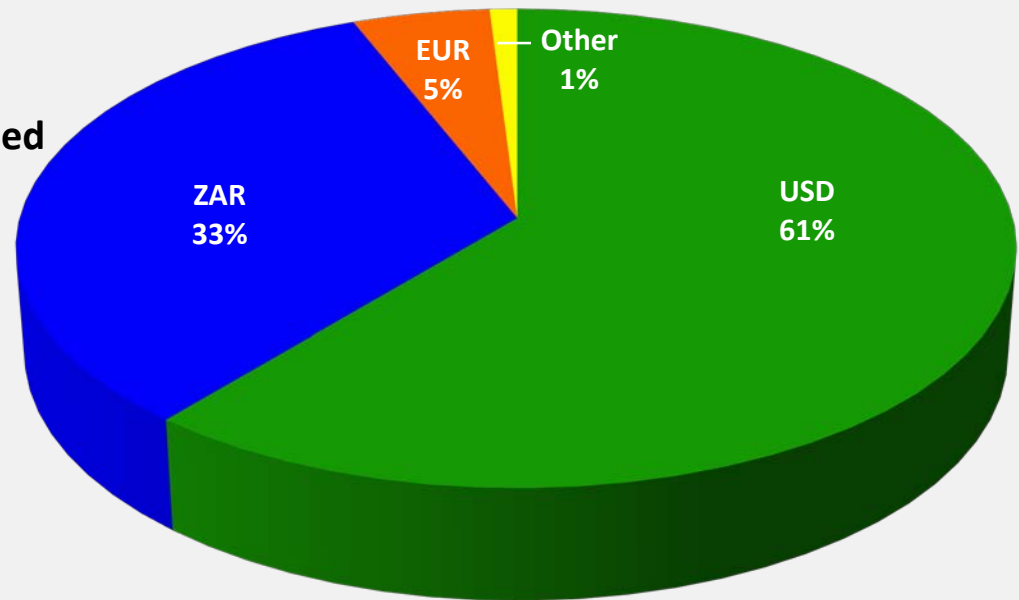
Allocation of Procurement Currencies

- Most API's and excipients are purchased mainly from Asia:

- ▶ India
- ▶ China

- Packaging materials are largely purchased from South African suppliers

- Aspen received an A-rating from Empowerdex in the preferential procurement category



Training and Development

- 140 new/ongoing 2010/11 Learnerships are in place, covering the fields of Business Administration, Equipment Setting and Operation, Safety Controls, Pharmaceutical Controls, etc.
- Additional Laboratory and Warehousing Learnerships are in development for 2011/2012
- 20 additional bursaries have been awarded towards tertiary studies in 2011
- 4 Pharmacists are receiving internships in 2011
- In 2010, 10 Apprentices as Fitters and 22 learners as Pharmacist Assistants were qualified



ARTISAN APPRENTICESHIP



PMA LEARNERSHIP



ABET

Environmental Sustainability Initiatives

Carbon Footprint

- Sophisticated air handling systems are in place to purify and filter the air discharged from the production areas
- Sampling was done in Port Elizabeth and East London for Sulphur Dioxide, Nitrous Oxide, Volatile Organic Compounds and particulate emissions: quantities were largely undetectable, or else, negligible
- Aspen is participating in a Carbon Disclosure Project, reporting using internationally accepted greenhouse gas accounting and reporting standards

Water Conservation

- The Port Elizabeth facility is currently re-using Reverse Osmosis reject water to feed its cooling towers and ablution facilities, coupled with switching off rotoclones when not in use, the project has yielded an average of 8% reduction in monthly water consumption
- East London is reusing cooling water, with a saving of 4% in monthly consumption

Environmental Sustainability Initiatives

Conservation of Electricity

- A target of 4% reduction in monthly energy consumption has been achieved through the installation of energy saving lighting in Liquids and Solids packing areas, and installation of lighting-off motion detectors across Port Elizabeth Administration Office Sites
- East London has achieved a 1% reduction in electricity consumption through installation of energy saving lights in the warehouses
- A feasibility study is in progress to evaluate the possibilities of generating and making use of alternative renewable energy sources



Social Sustainability Initiatives

- We have secured job security for our permanent employees despite recessionary pressures facing manufacturers in South Africa
- No permanent employees were made redundant by the implemented continuous improvement initiatives
- The Disabling Frequency Ratio of 0.63 against a tolerance of 1.00 demonstrates the effectiveness of safety standards at the facility
- More than 950 employees participated in the 2010 voluntary HIV/AIDS testing and counselling programme
 - ▶ 5% were HIV positive and are receiving the necessary treatment and support
- In addition to extensive on-the-job procedural and technical training that all staff receive in all aspects of operation, including GMP, SOPs and equipment operation, more than 10% of all Operations employees received external, specialised skills training in the six months to December 2010



Risks and Challenges

- **Realisation of awarded public sector volumes**

Mitigation: *International volumes are being introduced to Port Elizabeth*

- **Impact of inflation and currency movements on cost of goods**

Mitigation: *Benefits from continuous improvement initiatives and effective procurement strategies are being realised*

- **Shortage of scarce skills – pharmacists and artisan**

Mitigation: *Pharmacists retention programmes are in place, and investment is made in learnership programmes for assistance pharmacists and apprenticeships for artisans*

- **Increased competition from foreign suppliers**

Mitigation:

- ▶ *Cost of goods are benchmarked against international prices*
- ▶ *Aspen's successful performance in the recent ARV tender demonstrates competitiveness*
- ▶ *Aspen has proven to be a reliable supplier*
- ▶ *Significant investment has been made in creating flexible and diverse manufacturing capability to respond to current and future requirements*

Generics Bulletin Report : May 2010

